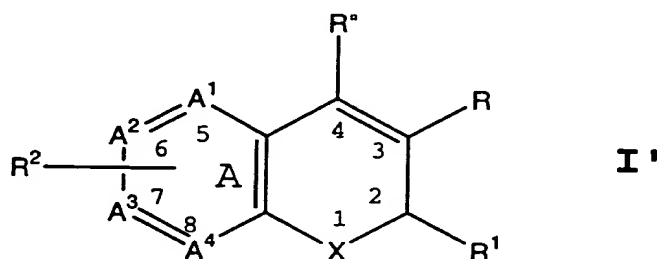


What is claimed is:

1. A compound of Formula I'



wherein X is selected from O, S, CR^cR^b and NR^a;
 wherein R^a is selected from hydrido, C₁-C₃-alkyl,
 (optionally substituted phenyl)-C₁-C₃-alkyl, acyl
 and carboxy-C₁-C₆-alkyl;

wherein each of R^b and R^c is independently selected
 from hydrido, C₁-C₃-alkyl, phenyl-C₁-C₃-alkyl, C₁-
 C₃-perfluoroalkyl, chloro, C₁-C₆-alkylthio, C₁-C₆-
 alkoxy, nitro, cyano and cyano-C₁-C₃-alkyl;

wherein R is selected from carboxyl, aminocarbonyl,
 C₁-C₆-alkylsulfonylaminocarbonyl and C₁-C₆-
 alkoxy carbonyl;

wherein R^o is selected from hydrido, phenyl, thienyl
 and C₂-C₆-alkenyl;

wherein Rⁱ is selected from C₁-C₃-perfluoroalkyl,
 chloro, C₁-C₆-alkylthio, C₁-C₆-alkoxy, nitro,
 cyano and cyano-C₁-C₃-alkyl;

wherein R² is one or more radicals independently
 selected from hydrido, halo, C₁-C₆-alkyl, C₂-C₆-
 alkenyl, C₂-C₆-alkynyl, halo-C₂-C₆-alkynyl, aryl-
 C₁-C₃-alkyl, aryl-C₂-C₆-alkynyl, aryl-C₂-C₆-
 alkenyl, C₁-C₆-alkoxy, methylenedioxy, C₁-C₆-
 alkylthio, C₁-C₆-alkylsulfinyl, aryloxy,
 arylthio, arylsulfinyl, heteroaryloxy, C₁-C₆-
 alkoxy-C₁-C₆-alkyl, aryl-C₁-C₆-alkyloxy,
 heteroaryl-C₁-C₆-alkyloxy, aryl-C₁-C₆-alkoxy-C₁-C₆-
 alkyl, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-
 haloalkylthio, C₁-C₆-haloalkylsulfinyl, C₁-C₆-

haloalkylsulfonyl, C₁-C₃-(haloalkyl-C₁-C₃-hydroxyalkyl, C₁-C₆-hydroxyalkyl, hydroxyimino-C₁-C₆-alkyl, C₁-C₆-alkylamino, arylamino, aryl-C₁-C₆-alkylamino, heteroaryl-amino, heteroaryl-C₁-C₆-alkylamino, nitro, cyano, amino, aminosulfonyl, C₁-C₆-alkylaminosulfonyl, arylaminosulfonyl, heteroarylaminosulfonyl, aryl-C₁-C₆-alkylaminosulfonyl, heteroaryl-C₁-C₆-alkylaminosulfonyl, heterocyclylsulfonyl, C₁-C₆-alkylsulfonyl, aryl-C₁-C₆-alkylsulfonyl, optionally substituted aryl, optionally substituted heteroaryl, aryl-C₁-C₆-alkylcarbonyl, heteroaryl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, arylcarbonyl, aminocarbonyl, C₁-C₆-alkoxycarbonyl, formyl, C₁-C₆-haloalkylcarbonyl and C₁-C₆-alkylcarbonyl; and wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least two of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a radical selected from naphthyl, quinolyl, isoquinolyl, quinoliziny, quinoxaliny and dibenzofuryl; or an isomer or pharmaceutically acceptable salt thereof.

2. A compound of Claim 1, wherein X is selected from O, S, CR^aR^b and NR^a; wherein R^a is selected from hydrido, C₁-C₃-alkyl, (optionally substituted phenyl)-C₁-C₃-alkyl, acyl and carboxy-C₁-C₆-alkyl; wherein each of R^b and R^b is independently selected from hydrido, C₁-C₃-alkyl, phenyl-C₁-C₃-alkyl, C₁-C₃-perfluoroalkyl, chloro, C₁-C₆-alkylthio, C₁-C₆-alkoxy, nitro, cyano and cyano-C₁-C₃-alkyl; wherein R is selected from carboxyl, aminocarbonyl, C₁-C₆-alkylsulfonylaminocarbonyl and C₁-C₆-alkoxycarbonyl; wherein R^a is selected from hydrido, phenyl, thienyl and C₂-C₆-alkenyl; wherein R¹ is selected from C₁-C₃-

09496695-020200

perfluoroalkyl, chloro, C₁-C₆-alkylthio, C₁-C₆-alkoxy, nitro, cyano and cyano-C₁-C₃-alkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl, halo-C₂-C₆-alkynyl, aryl-C₁-C₃-alkyl, aryl-C₂-C₆-alkynyl, aryl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, methylenedioxy, C₁-C₆-alkylthio, C₁-C₆-alkylsulfinyl, aryloxy, arylthio, arylsulfinyl, heteroaryloxy, C₁-C₆-alkoxy-C₁-C₆-alkyl, aryl-C₁-C₆-alkyloxy, heteroaryl-C₁-C₆-alkyloxy, aryl-C₁-C₆-alkoxy-C₁-C₆-alkyl, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-haloalkylthio, C₁-C₆-haloalkylsulfinyl, C₁-C₆-haloalkylsulfonyl, C₁-C₃-(haloalkyl-C₁-C₃-hydroxyalkyl, C₁-C₆-hydroxyalkyl, hydroxyimino-C₁-C₆-alkyl, C₁-C₆-alkylamino, arylamino, aryl-C₁-C₆-alkylamino, heteroarylamino, heteroaryl-C₁-C₆-alkylamino, nitro, cyano, amino, aminosulfonyl, C₁-C₆-alkylaminosulfonyl, arylaminosulfonyl, heteroarylaminosulfonyl, aryl-C₁-C₆-alkylaminosulfonyl, heteroaryl-C₁-C₆-alkylaminosulfonyl, heterocyclylsulfonyl, C₁-C₆-alkylsulfonyl, aryl-C₁-C₆-alkylsulfonyl, optionally substituted aryl, optionally substituted heteroaryl, aryl-C₁-C₆-alkylcarbonyl, heteroaryl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, arylcarbonyl, aminocarbonyl, C₁-C₆-alkoxycarbonyl, formyl, C₁-C₆-haloalkylcarbonyl and C₁-C₆-alkylcarbonyl; and wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

3. A compound of Claim 2 wherein X is selected from O, S and NR^a; wherein R^a is selected from hydrido, C₁-C₃-alkyl and (optionally substituted phenyl)methyl; wherein R is carboxyl; wherein R^b is selected from hydrido and C₂-C₆-alkenyl; wherein R¹ is

002020 5696460

selected from C₁-C₃-perfluoroalkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl, halo-C₂-C₆-alkynyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl, phenyl-C₂-C₆-alkenyl, C₁-C₃-alkoxy, methylenedioxy, C₁-C₃-alkoxy-C₁-C₃-alkyl, C₁-C₃-alkylthio, C₁-C₃-alkylsulfinyl, phenyloxy, phenylthio, phenylsulfinyl, C₁-C₃-haloalkyl-C₁-C₃-hydroxyalkyl, phenyl-C₁-C₃-alkyloxy-C₁-C₃-alkyl, C₁-C₃-haloalkyl, C₁-C₃-haloalkoxy, C₁-C₃-haloalkylthio, C₁-C₃-hydroxyalkyl, C₁-C₃-alkoxy-C₁-C₃-alkyl, hydroxyimino-C₁-C₃-alkyl, C₁-C₆-alkylamino, nitro, cyano, amino, aminosulfonyl, N-alkylaminosulfonyl, N-arylaminosulfonyl, N-heteroarylaminosulfonyl, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, phenyl-C₁-C₃-alkylsulfonyl, 5- to 8-membered heterocyclylsulfonyl, C₁-C₆-alkylsulfonyl, optionally substituted phenyl, optionally substituted 5- to 9-membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl, phenylcarbonyl, 4-chlorophenylcarbonyl, 4-hydroxyphenylcarbonyl, 4-trifluoromethylphenylcarbonyl, 4-methoxyphenylcarbonyl, aminocarbonyl, formyl, and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl, benzofurylphenyl, or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

4. A compound of Claim 3 wherein X is selected from O, S and NR^a; wherein R^a is selected from hydrido, methyl, ethyl, (4-trifluoromethyl)benzyl, (4-chloromethyl)benzyl, (4-methoxy)benzyl, and (4-cyano)benzyl, (4-nitro)benzyl; wherein R is carboxyl; wherein R^b is selected from hydrido and ethenyl; wherein R¹ is selected from trifluoromethyl

and pentafluoroethyl; wherein R² is one or more radicals independently selected from hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenylethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy, methylthio, methylsulfinyl, phenyloxy, phenylthio, phenylsulfinyl, methylenedioxy, benzyloxymethyl, trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl, hydroxy-trifluoroethyl, methoxymethyl, hydroxyiminomethyl, N-methylamino, nitro, cyano, amino, aminosulfonyl, N-methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, optionally substituted phenylcarbonyl, aminocarbonyl, formyl and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl, or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

5. A compound of Claim 4 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of 6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

- 7-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 2,7-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 7-bromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 6-chloro-7-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6-chloro-8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-ethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 8-bromo-6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-trifluoromethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 8-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 35 7-isopropoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

002020-569646

- 8-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7,8-dimethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6,8-bis(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 8-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 6,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6,8-dibromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dimethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-nitro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-amino-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- ethyl 6-amino-2-trifluoromethyl-2H-1-benzopyran-3-carboxylate;
- 35 6-chloro-8-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

002020" 5696460

- 8-chloro-6-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6,8-difluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
5 6-bromo-8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
8-bromo-6-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
8-bromo-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
10 8-bromo-5-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-chloro-8-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
15 6-bromo-8-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
7-(N,N-diethylamino)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-[[(phenylmethyl) amino] sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
20 6-[(dimethylamino) sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-aminosulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
25 6-(methylamino) sulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-[(4-morpholino) sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-[(1,1-dimethylethyl) aminosulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
30 6-[(2-methylpropyl) aminosulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-methylsulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
35 8-chloro-6-[[(phenylmethyl) amino] sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

002020" 5696460

- 6-N,N-diethylaminosulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-phenylacetyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6-(2,2-dimethylpropylcarbonyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-7-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid;
- 10 6-[[(2-furanylmethyl) amino] sulfonyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-[(phenylmethyl) sulfonyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 15 6-[[(phenylethyl) amino] sulfonyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-iodo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 6-chloro-8-iodo-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 8-bromo-6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6-formyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-formyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-7-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 30 5,6-dichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid ;
- 35 6-hydroxymethyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(difluoromethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;

002020" 56996460

- 2,6-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5,6,7-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5 6,7,8-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(methylthio)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(methylsulfinyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 10 5,8-dichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(pentafluoroethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 15 6-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-7-methyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 20 6-chloro-2,7-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5-methoxy-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 25 6-benzoyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-chlorobenzoyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-hydroxybenzoyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 30 6-phenoxy-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-6-(4-chlorophenoxy)-2-trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 35 2-(trifluoromethyl)-6-[4-(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-methoxyphenoxy)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;

002020"5699460

- 6-(3-chloro-4-methoxyphenoxy)-2-(trifluoromethyl)-
2H-1-benzopyran-3-carboxylic acid;
6-(4-chlorophenoxy)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
5 8-chloro-2-(trifluoromethyl)-6-[4-
(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-
carboxylic acid;
6-chloro-8-cyano-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
10 6-chloro-8-[(hydroxyimino)methyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
6-chloro-8-(hydroxymethyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
15 8-(1H-benzimidazol-2-yl)-6-chloro-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
7-(1,1-dimethylethyl)-2-(pentafluoroethyl)-2H-1-
benzopyran-3-carboxylic acid;
20 6-chloro-8-(methoxymethyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
6-chloro-8-(benzyloxymethyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
25 6-chloro-8-ethenyl-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
6-chloro-8-ethynyl-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
6-chloro-8-(2-thienyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
30 6-chloro-8-(2-furanyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
6-chloro-8-(5-chloro-1-pentynyl)-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
35 6-chloro-8-(1-pentynyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
6-chloro-8-(phenylethynyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

0049695-02000

- 6-chloro-8-(3,3-dimethyl-1-butynyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5 6-chloro-8-[(4-chlorophenyl)ethynyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-[(4-methoxyphenyl)ethynyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 10 6-(phenylethynyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(4-chlorophenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(3-methoxyphenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 15 6-chloro-8-[(4-methylthio)phenyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-[(4-methylsulfonyl)phenyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 20 6-chloro-8-phenyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-8-fluoro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 25 6-(4-fluorophenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-phenyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 30 8-chloro-6-fluoro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6,8-diiodo-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(5-chloro-2-thienyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 35 6-(2-thienyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;

046655-0000

- 6-(4-chlorophenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-bromophenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5 6-(ethynyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-methyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(4-methoxyphenyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 6-chloro-2-(trifluoromethyl)-4-ethenyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-2-(trifluoromethyl)-4-phenyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6-chloro-4-(2-thienyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(2,2,2-trifluoro-1-hydroxyethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 20 6-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 6,8-dimethyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 25 6-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 6,7-dimethyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 30 8-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 35 6-chloro-7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
- 7-chloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

002020-5695460

- 6,7-dichloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-benzopyran-3-carboxylic acid;
- 5 6,8-dichloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid;
6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6,8-dichloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 10 6,7-difluoro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-iodo-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 15 6-bromo-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
1,2-dihydro-6-(trifluoromethoxy)-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-(trifluoromethyl)-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 20 6-cyano-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-1-methyl-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 25 6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[[4-(trifluoromethyl)phenyl]methyl]-3-quinolinecarboxylic acid;
6-chloro-1-[(4-chlorophenyl)methyl]-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 30 6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[[4-(methoxy)phenyl]methyl]-3-quinolinecarboxylic acid;
6-chloro-1-[(4-cyanophenyl)methyl]-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 35 6-chloro-1,2-dihydro-1-[(4-nitrophenyl)methyl]-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-1-ethyl-2-(trifluoromethyl)-3-quinolinecarboxylic acid;

002020 "5696460

- 6-chloro-2-(trifluoromethyl)-1,2-dihydro[1,8]naphthyridine-3-carboxylic acid;
 2-trifluoromethyl-2H-naphtho[1,2-b]pyran-3-carboxylic acid;
 5 2-trifluoromethyl-3H-naphtho[2,1-b]pyran-3-carboxylic acid;
 2-trifluoromethyl-2H-naphtho[2,3-b]pyran-3-carboxylic acid;
 5-(hydroxymethyl)-8-methyl-2-(trifluoromethyl)-2H-pyrano[2,3-c]pyridine-3-carboxylic acid;
 10 6-(trifluoromethyl)-6h-1,3-dioxolo[4,5-g][1]benzopyran-7-carboxylic acid; and
 3-(trifluoromethyl)-3H-benzofuro[3,2-f][1]benzopyran-2-carboxylic acid.

- 15 6. A compound of Claim 2 wherein X is O; wherein R is carboxyl; wherein R' is selected from hydrido and C₂-C₆-alkenyl; wherein R¹ is selected from C₁-C₃-perfluoroalkyl; wherein R² is one or more
 20 radicals independently selected from hydrido, halo, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl, phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, phenyloxy, 5- or 6-membered heteroaryloxy, phenyl-C₁-C₆-alkyloxy, 5- or 6-membered heteroaryl-C₁-C₆-alkyloxy, C₁-C₆-
 25 haloalkyl, C₁-C₆-haloalkoxy, N-(C₁-C₆-alkyl)amino, N,N-di-(C₁-C₆-alkyl)amino, N-phenylamino, N-(phenyl-C₁-C₆-alkyl)amino, N-heteroarylamino, N-(heteroaryl-C₁-C₆-alkyl)amino, nitro, amino, aminosulfonyl, N-(C₁-C₆-alkyl)aminosulfonyl, N,N-di-(C₁-C₆-
 30 alkyl)aminosulfonyl, N-arylaminosulfonyl, N-heteroarylaminosulfonyl, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, 5- to 8-membered heterocyclylsulfonyl, C₁-C₆-alkylsulfonyl, optionally
 35 substituted phenyl, optionally substituted 5- or 6-membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, phenylcarbonyl, aminocarbonyl,

002020-5696460

- and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or an isomer or
- 5 pharmaceutically acceptable salt thereof.

7. A compound of Claim 6 wherein X is O; wherein R is carboxyl; wherein R' is selected from hydrido and ethenyl; wherein R¹ is selected from
- 10 trifluoromethyl and pentafluoroethyl; wherein R² is one or more radicals independently selected from hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl,
- 15 phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy, methylthio, methylsulfinyl, phenyloxy, phenylthio, phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy, phenylmethoxy, methylenedioxy, benzyloxymethyl,
- 20 trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl, hydroxy-trifluoroethyl, methoxymethyl, hydroxyiminomethyl, N-methylamino, N-phenylamino, N-(benzyl)amino, nitro, cyano, amino,
- 25 aminosulfonyl, N-methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl
- 30 substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl,
- 35 phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; and wherein one of the A ring atoms A¹, A², A³ and A⁴ is nitrogen and the other three are

002020" 56996460

carbon; or an isomer or pharmaceutically acceptable salt thereof.

9. A compound of Claim 7 wherein X is O;
- 5 wherein R is carboxyl; wherein R" is selected from hydrido and ethenyl; wherein R¹ is selected from trifluoromethyl and pentafluoroethyl; wherein R² is one or more radicals independently selected from
- 10 hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy, methylthio, methylsulfinyl, phenyloxy, phenylthio,
- 15 phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy, phenylmethoxy, methylenedioxy, benzyloxymethyl, trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl, hydroxy-trifluoroethyl,
- 20 methoxymethyl, hydroxyiminomethyl, N-methylamino, N-phenylamino, N-(benzyl)amino, nitro, cyano, amino, aminosulfonyl, N-methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl,
- 25 benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl
- 30 substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl, phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically
- 35 acceptable salt thereof.

0022020"5696460

10. A compound of Claim 9 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of
- 5 6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
(S)-6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-chloro-7-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
(S)-6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6-chloro-8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6-trifluoromethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 (S)-6-trifluoromethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
(S)-6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
6,8-dichloro-7-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid;
(S)-6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid;
- 35 6-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
(S)-6-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;

002020" 5696460

- 6-hydroxymethyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(difluoromethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5 2,6-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5,6,7-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6,7,8-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 10 6-(methylthio)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(pentafluoroethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 15 2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-7-methyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-benzoyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 20 6-(4-chlorobenzoyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-hydroxybenzoyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 25 6-phenoxy-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 2-(trifluoromethyl)-6-[4-(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- (S)-2-(trifluoromethyl)-6-[4-(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- 30 6-(4-methoxyphenoxy)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(3-chloro-4-methoxyphenoxy)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 35 6-(4-chlorophenoxy)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;

002020"5695460

- 8-chloro-2-(trifluoromethyl)-6-[4-(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- 5 6-chloro-8-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(2-thienyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(phenylethynyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 10 6-chloro-8-[(4-chlorophenyl)ethynyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-[(4-methoxyphenyl)ethynyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 15 (S)-6-chloro-8-[(4-methoxyphenyl)ethynyl]-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(phenylethynyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 20 6-chloro-8-(4-chlorophenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-phenyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 25 6-(4-bromophenyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(4-methoxyphenyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid; and
- 6-(2,2,2-trifluoro-1-hydroxyethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid.
- 30

11. A compound of Claim 2 wherein X is S; wherein R is carboxyl; wherein R¹ is selected from
- 35 C₁-C₆-perfluoroalkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl, phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, phenyloxy, 5- or

0022020" 5696460

6-membered heteroaryloxy, phenyl-C₁-C₆-alkyloxy, 5-
 or 6-membered heteroaryl-C₁-C₆-alkyloxy, C₁-C₆-
 haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-alkylamino, N-
 phenylamino, N-(phenyl-C₁-C₆-alkyl)amino, N-
 5 heteroarylamino, N-(heteroaryl-C₁-C₆-alkylamino,
 nitro, amino, aminosulfonyl, N-alkylaminosulfonyl,
 N-arylaminosulfonyl, N-heteroarylamino, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-
 C₆-alkyl)aminosulfonyl, 5- to 8-membered
 10 heterocyclylsulfonyl, C₁-C₆-alkylsulfonyl, optionally
 substituted phenyl, optionally substituted 5- or 6-
 membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl,
 heteroarylcarbonyl, phenylcarbonyl, aminocarbonyl,
 and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹,
 15 A², A³ and A⁴ are independently selected from carbon
 and nitrogen with the proviso that at least three of
 A¹, A², A³ and A⁴ are carbon; or an isomer or
 pharmaceutically acceptable salt thereof.

20 12. A compound of Claim 11 wherein X is S;
 wherein R is carboxyl; wherein R' is selected from
 hydrido and ethenyl; wherein R¹ is selected from
 trifluoromethyl and pentafluoroethyl; wherein R² is
 one or more radicals independently selected from
 25 hydrido, chloro, bromo, fluoro, iodo, methyl, tert-
 butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-
 pentynyl, 3,3-dimethyl-1-butynyl, benzyl,
 phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl,
 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy,
 30 methylthio, methylsulfinyl, phenyloxy, phenylthio,
 phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy,
 phenylmethoxy, methylenedioxy, benzyloxymethyl,
 trifluoromethyl, difluoromethyl, pentafluoroethyl,
 trifluoromethoxy, trifluoromethylthio,
 35 hydroxymethyl, hydroxy-trifluoroethyl,
 methoxymethyl, hydroxyiminomethyl, N-methylamino, N-
 phenylamino, N-(benzyl)amino, nitro, cyano, amino,
 aminosulfonyl, N-methylaminosulfonyl, N-

002020"5596460

phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl, phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically acceptable salt thereof.

13. A compound of Claim 12 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of

6-chloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

6-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

6,8-dimethyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

6-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

6,7-dimethyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

8-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

6-chloro-7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

7-chloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

- 6,7-dichloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
 2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-benzopyran-3-carboxylic acid; and
 5 6,8-dichloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid.

14. A compound of Claim 2 wherein X is NR^a; wherein R^a is selected from hydrido, C₁-C₃-alkyl, phenyl-C₁-C₃-alkyl, acyl and carboxy-C₁-C₃-alkyl; wherein R is carboxyl; wherein R¹ is selected from C₁-C₃-perfluoroalkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl,
 15 phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, phenyloxy, 5- or 6-membered heteroaryloxy, phenyl-C₁-C₆-alkyloxy, 5- or 6-membered heteroaryl-C₁-C₆-alkyloxy, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-alkylamino, N-phenylamino, N-(phenyl-C₁-C₆-alkyl)amino, N-heteroarylamino, N-
 20 (heteroaryl-C₁-C₆-alkylamino, nitro, amino, aminosulfonyl, N-alkylaminosulfonyl, N-arylaminosulfonyl, N-heteroarylamino, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, 5- to 8-membered
 25 heterocyclisulfonyl, C₁-C₆-alkylsulfonyl, optionally substituted phenyl, optionally substituted 5- or 6-membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, phenylcarbonyl, aminocarbonyl, and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically acceptable salt thereof.

- 35 15. A compound of Claim 14 wherein X is NR^a; wherein R^a is selected from hydrido, methyl, ethyl, (4-trifluoromethyl)benzyl, (4-chloromethyl)benzyl, (4-methoxy)benzyl, (4-cyano)benzyl, and (4-

00220"599665-020200

nitro)benzyl; wherein R is carboxyl; wherein R" is selected from hydrido and ethenyl; wherein R¹ is selected from trifluoromethyl and pentafluoroethyl; wherein R² is one or more radicals independently
5 selected from hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy,
10 methylthio, methylsulfinyl, phenyloxy, phenylthio, phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy, phenylmethoxy, methylenedioxy, benzyloxymethyl, trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl,
15 hydroxy-trifluoroethyl, methoxymethyl, hydroxyiminomethyl, N-methylamino, N-phenylamino, N-(benzyl)amino, nitro, cyano, amino, aminosulfonyl, N-methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, benzylsulfonyl,
20 phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl,
25 benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl, phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are carbon; or an
30 isomer or pharmaceutically acceptable salt thereof.

16. A compound of Claim/15 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of

35 6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;

- 6,8-dichloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6,7-difluoro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
5 6-iodo-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-bromo-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
1,2-dihydro-6-(trifluoromethoxy)-2-(trifluoromethyl)-
10 3-quinolinecarboxylic acid;
6-(trifluoromethyl)-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-cyano-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
15 6-chloro-1,2-dihydro-1-methyl-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[[4-(trifluoromethyl)phenyl]methyl]-3-quinolinecarboxylic acid;
20 6-chloro-1-[(4-chlorophenyl)methyl]-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[[4-(methoxy)phenyl]methyl]-3-quinolinecarboxylic acid;
25 6-chloro-1-[(4-cyanophenyl)methyl]-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-1-[(4-nitrophenyl)methyl]-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-1-ethyl-2-(trifluoromethyl)-3-
30 quinolinecarboxylic acid; and
(S)-6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid.

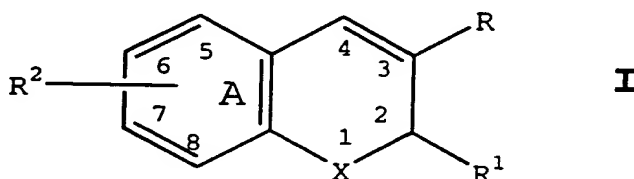
17. A compound of Claim 2 wherein X is selected
35 from O, S and NR^a ; wherein R^a is selected from hydrido, $\text{C}_1\text{-C}_3$ -alkyl, phenyl- $\text{C}_1\text{-C}_3$ -alkyl, acyl and carboxy- $\text{C}_1\text{-C}_3$ -alkyl; wherein R is selected from carboxyl; wherein R^1 is selected from $\text{C}_1\text{-C}_3$ -

perfluoroalkyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; and wherein R² together with ring A
5 forms a naphthyl or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

18. A compound of Claim 17 wherein X is selected from O, S and NR^a; wherein R^a is selected from
10 hydrido, methyl, ethyl, (4-trifluoromethyl)benzyl, (4-chloromethyl)benzyl, (4-methoxy)benzyl, and (4-cyano)benzyl, (4-nitro)benzyl; wherein R is carboxyl; wherein R^b is selected from hydrido and ethenyl; wherein R^c is selected from trifluoromethyl and
15 pentafluoroethyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl, or quinolyl radical; or an
20 isomer or pharmaceutically acceptable salt thereof.

19. A compound of Claim 18 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of
25 2-trifluoromethyl-2H-naphtho[1,2-b]pyran-3-carboxylic acid;
2-trifluoromethyl-3H-naphtho[2,1-b]pyran-3-carboxylic acid;
30 2-trifluoromethyl-2H-naphtho[2,3-b]pyran-3-carboxylic acid;
5-(hydroxymethyl)-8-methyl-2-(trifluoromethyl)-2H-pyrano[2,3-c]pyridine-3-carboxylic acid;
6-(trifluoromethyl)-6h-1,3-dioxolo[4,5-
35 g][1]benzopyran-7-carboxylic acid; and
3-(trifluoromethyl)-3H-benzofuro[3,2-f][1]benzopyran-2-carboxylic acid.

20. A compound of Formula I



5

wherein X is selected from O or S or NR^a;

wherein R^a is alkyl;

wherein R is selected from carboxyl, aminocarbonyl, alkylsulfonylaminocarbonyl and alkoxy carbonyl;

10

wherein R¹ is selected from haloalkyl, alkyl, aralkyl, cycloalkyl and aryl optionally substituted with one or more radicals selected from alkylthio, nitro and alkylsulfonyl; and

15

wherein R² is one or more radicals selected from hydrido, halo, alkyl, aralkyl, alkoxy, aryloxy, heteroaryloxy, aralkyloxy, heteroaralkyloxy, haloalkyl, haloalkoxy, alkylamino, arylamino, aralkylamino, heteroaryl amino, heteroarylalkyl amino, nitro, amino, aminosulfonyl, alkylaminosulfonyl, arylaminosulfonyl, heteroarylaminosulfonyl, aralkylaminosulfonyl, heteroaralkylaminosulfonyl, heterocyclosulfonyl, alkylsulfonyl, optionally substituted aryl, optionally substituted heteroaryl, aralkylcarbonyl, heteroarylcarbonyl, arylcarbonyl, aminocarbonyl, and alkylcarbonyl;

20

25

or wherein R² together with ring A forms a naphthyl radical;

or an isomer or pharmaceutically acceptable salt thereof.

30

21. Compound of Claim 20 wherein X is oxygen or sulfur; wherein R is selected from carboxyl, lower alkyl, lower aralkyl and lower alkoxy carbonyl;

wherein R¹ is selected from lower haloalkyl, lower cycloalkyl and phenyl; and wherein R² is one or more radicals selected from hydrido, halo, lower alkyl, lower alkoxy, lower haloalkyl, lower haloalkoxy, lower alkylamino, nitro, amino, aminosulfonyl, lower alkylaminosulfonyl, 5- or 6- membered heteroarylalkylaminosulfonyl, lower aralkylaminosulfonyl, 5- or 6- membered nitrogen containing heterocyclosulfonyl, lower alkylsulfonyl, optionally substituted phenyl, lower aralkylcarbonyl, and lower alkylcarbonyl; or wherein R² together with ring A forms a naphthyl radical; or an isomer or pharmaceutically acceptable salt thereof.

22. Compound of Claim 21 wherein X is oxygen or sulfur; wherein R is carboxyl; wherein R¹ is lower haloalkyl; and wherein R² is one or more radicals selected from hydrido, halo, lower alkyl, lower haloalkyl, lower haloalkoxy, lower alkylamino, amino, aminosulfonyl, lower alkylaminosulfonyl, 5- or 6- membered heteroarylalkylaminosulfonyl, lower aralkylaminosulfonyl, lower alkylsulfonyl, 6- membered nitrogen containing heterocyclosulfonyl, optionally substituted phenyl, lower aralkylcarbonyl, and lower alkylcarbonyl; or wherein R² together with ring A forms a naphthyl radical; or an isomer or pharmaceutically acceptable salt thereof.

23. Compound of Claim 22 wherein R is carboxyl; wherein R¹ is selected from fluoromethyl, chloromethyl, dichloromethyl, trichloromethyl, pentafluoroethyl, heptafluoropropyl, difluoroethyl, difluoropropyl, dichloroethyl, dichloropropyl, difluoromethyl, and trifluoromethyl; and wherein R² is one or more radicals selected from hydrido, chloro, fluoro, bromo, iodo, methyl, ethyl, isopropyl, tert-butyl, butyl, isobutyl, pentyl, hexyl, methoxy, ethoxy, isopropoxy, tertbutoxy, trifluoromethyl,

- 002020" 55996460
- difluoromethyl, trifluoromethoxy, amino, N,N-dimethylamino, N,N-diethylamino, N-phenylmethylenesulfonyl, N-phenylethylmethylenesulfonyl, N-(2-furylmethyl)aminosulfonyl, nitro, N,N-dimethylaminosulfonyl, aminosulfonyl, N-methylaminosulfonyl, N-ethylsulfonyl, 2,2-dimethylethylaminosulfonyl, N,N-dimethylaminosulfonyl, N-(2-methylpropyl)aminosulfonyl, N-morpholinosulfonyl, methylsulfonyl, benzylcarbonyl, 2,2-dimethylpropylcarbonyl, phenylacetyl and phenyl; or wherein R² together with ring A forms a naphthyl radical; or an isomer or pharmaceutically acceptable salt thereof.

24. Compound of Claim 23 wherein R is carboxyl; wherein R¹ is trifluoromethyl or pentafluorethyl; and wherein R² is one or more radicals selected from
- hydrido, chloro, fluoro, bromo, iodo, methyl, ethyl, isopropyl, tert-butyl, methoxy, trifluoromethyl, trifluoromethoxy, N-phenylmethylenesulfonyl, N-phenylethylmethylenesulfonyl, N-(2-furylmethyl)aminosulfonyl, N,N-dimethylaminosulfonyl, N-methylaminosulfonyl, N-(2,2-dimethylethyl)aminosulfonyl, dimethylaminosulfonyl, 2-methylpropylaminosulfonyl, N-morpholinosulfonyl, methylsulfonyl, benzylcarbonyl, and phenyl; or wherein R² together with ring A forms a naphthyl radical; or an isomer or pharmaceutically acceptable salt thereof.

25. A compound of Claim 24 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of
- 6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

- 6-chloro-7-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 2-trifluoromethyl-3H-naphthopyran-3-carboxylic acid ;
- 10 7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6-trifluoromethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 8-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7,8-dimethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-bis(dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 7-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-chloro-7-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

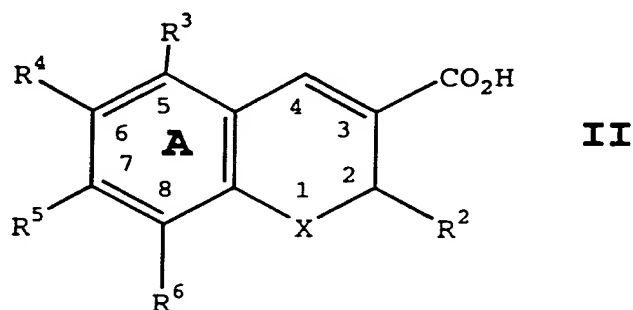
002020"55996460

- 6-chloro-8-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 2-trifluoromethyl-3H-naptho[2,1-b]pyran-3-carboxylic acid;
- 6-chloro-8-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 8-chloro-6-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-bromo-6-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 8-bromo-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-bromo-5-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6-chloro-8-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-8-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[[(phenylmethyl) amino] sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-[(dimethylamino) sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

002020"5696460

- 6-[(methylamino)sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[(4-morpholino)sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6-[(1,1-dimethylethyl)aminosulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[(2-methylpropyl)aminosulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-methylsulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 8-chloro-6-[[(phenylmethyl) amino] sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-phenylacetyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6,8-dibromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-5,6-dimethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-(S)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 6-benzylsulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[[N-(2-furylmethyl) amino] sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6-[[N-(2-phenylethyl) amino] sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-iodo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-(1,1-dimethylethyl)-2-pentafluoroethyl-2H-1-benzopyran-3-carboxylic acid; and
- 30 6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid.

26. A compound of Formula II



wherein X is O or S;

wherein R² is lower haloalkyl;

5 wherein R³ is selected from hydrido, and halo;

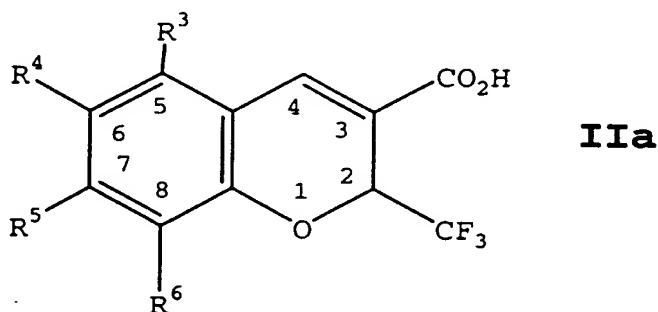
wherein R⁴ is selected from hydrido, halo, lower
alkyl, lower haloalkoxy, lower alkoxy, lower
aralkylcarbonyl, lower dialkylaminosulfonyl, lower
alkylaminosulfonyl, lower aralkylaminosulfonyl, lower
10 heteroaralkylaminosulfonyl, and 5- or 6- membered
nitrogen-containing heterocyclosulfonyl;

wherein R⁵ is selected from hydrido, lower alkyl,
halo, lower alkoxy, and aryl; and

wherein R⁶ is selected from hydrido, halo, lower
15 alkyl, lower alkoxy, and aryl;

or an isomer or pharmaceutically acceptable salt
thereof.

27. A compound of Formula IIa:

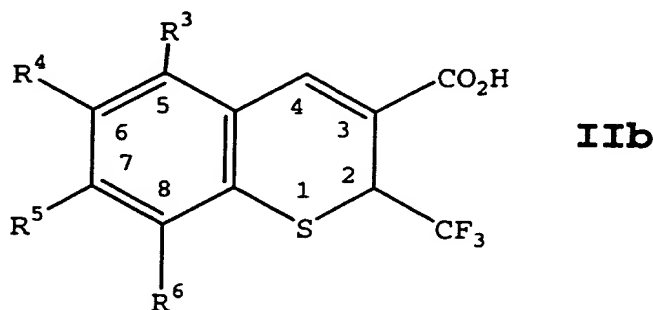


wherein R³ is selected from hydrido, lower alkyl,
lower hydroxyalkyl, lower alkoxy and halo;

- wherein R⁴ is selected from hydrido, halo, lower alkyl, lower alkylthio, lower haloalkyl, amino, aminosulfonyl, lower alkylsulfonyl, lower alkylsulfinyl, lower alkoxyalkyl, lower alkylcarbonyl, formyl, cyano, lower haloalkylthio, substituted or unsubstituted phenylcarbonyl, lower haloalkoxy, lower alkoxy, lower aralkylcarbonyl, lower dialkylaminosulfonyl, lower alkylaminosulfonyl, lower aralkylaminosulfonyl, lower heteroaralkylaminosulfonyl, 5- or 6- membered heteroaryl, lower hydrooxyalkyl, optionally substituted phenyl and 5- or 6- membered nitrogen containing heterocyclosulfonyl; wherein R⁵ is selected from hydrido, lower alkyl, halo, lower haloalkyl, lower alkoxy, and phenyl; and wherein R⁶ is selected from hydrido, halo, cyano, hydroxyiminomethyl, lower hydroxyalkyl, lower alkynyl, phenylalkynyl, lower alkyl, lower alkoxy, formyl and phenyl; or an isomer or pharmaceutically acceptable salt thereof.

28. Compound of Claim 27 wherein R¹ is selected from hydrido, and chloro; wherein R⁴ is selected from chloro, methyl, tert-butyl, methylthio, trifluoromethyl, difluoromethyl, pentafluoromethyl, trifluoromethylsulfide, trifluoromethoxy, cyano, substituted or unsubstituted phenylcarbonyl, and substituted or unsubstituted phenyl; wherein R⁵ is selected from hydrido, methyl, tert-butyl, chloro; and wherein R⁶ is selected from hydrido, chloro, thienyl, hydroxyiminomethyl, substituted or unsubstituted phenylethynyl, and substituted or unsubstituted phenyl; or an isomer or pharmaceutically acceptable salt thereof.

29. A compound of Formula IIb:



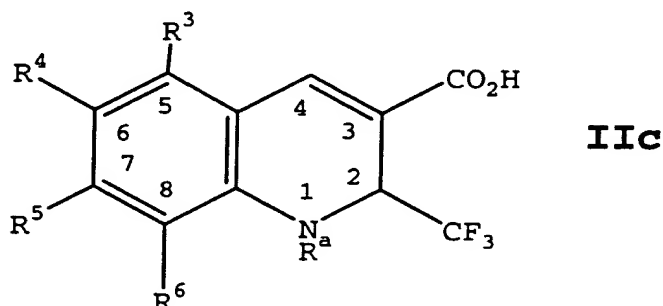
wherein R³ is selected from hydrido, lower alkyl,
 lower hydroxyalkyl, lower alkoxy and halo; wherein R⁴
 5 is selected from hydrido, halo, lower alkyl, lower
 alkylthio, lower haloalkyl, amino, aminosulfonyl,
 lower alkylsulfonyl, lower alkylsulfinyl, lower
 alkoxyalkyl, lower alkylcarbonyl, formyl, cyano,
 lower haloalkylthio, substituted or unsubstituted
 10 phenylcarbonyl, lower haloalkoxy, lower alkoxy, lower
 aralkylcarbonyl, lower dialkylaminosulfonyl, lower
 alkylaminosulfonyl, lower aralkylaminosulfonyl, lower
 heteroaralkylaminosulfonyl, 5- or 6- membered
 heteroaryl, lower hydroxyalkyl, optionally
 15 substituted phenyl and 5- or 6- membered nitrogen
 containing heterocyclosulfonyl; wherein R⁵ is selected
 from hydrido, lower alkyl, halo, lower haloalkyl,
 lower alkoxy, and phenyl; and wherein R⁶ is selected
 from hydrido, halo, cyano, hydroxyiminomethyl, lower
 20 hydroxyalkyl, lower alkynyl, phenylalkynyl, lower
 alkyl, lower alkoxy, formyl and phenyl; or an isomer
 or pharmaceutically acceptable salt thereof.

30. Compound of Claim 29 wherein R³ is selected
 25 from hydrido, and chloro; wherein R⁴ is selected from
 chloro, methyl, tert-butyl, methylthio,
 trifluoromethyl, difluoromethyl, pentafluoromethyl,
 trifluoromethylsulfide, trifluoromethoxy, cyano,
 substituted or unsubstituted phenylcarbonyl, and
 30 substituted or unsubstituted phenyl; wherein R⁵ is
 selected from hydrido, methyl, tert-butyl, chloro;

002020" 56996460

- and wherein R^6 is selected from hydrido, chloro, thienyl, hydroxyiminomethyl, substituted or unsubstituted phenylethynyl, and substituted or unsubstituted phenyl; or an isomer or
- 5 pharmaceutically acceptable salt thereof.

31. A compound of Formula IIc:



- 10 wherein R^a is selected from hydrido and lower aralkyl; wherein R^3 is selected from hydrido, lower alkyl, lower hydroxyalkyl, lower alkoxy and halo; wherein R^4 is selected from hydrido, halo, lower alkyl, lower
- 15 alkylthio, lower haloalkyl, amino, aminosulfonyl, lower alkylsulfonyl, lower alkylsulfinyl, lower alkoxyalkyl, lower alkylcarbonyl, formyl, cyano, lower haloalkylthio, substituted or unsubstituted phenylcarbonyl, lower haloalkoxy, lower alkoxy, lower
- 20 aralkylcarbonyl, lower dialkylaminosulfonyl, lower alkylaminosulfonyl, lower aralkylaminosulfonyl, lower heteroaralkylaminosulfonyl, 5- or 6- membered heteroaryl, lower hydroxyalkyl, optionally substituted phenyl and 5- or 6- membered nitrogen
- 25 containing heterocyclosulfonyl; wherein R^5 is selected from hydrido, lower alkyl, halo, lower haloalkyl, lower alkoxy, and phenyl; and wherein R^6 is selected from hydrido, halo, cyano, hydroxyiminomethyl, lower hydroxyalkyl, lower alkynyl, phenylalkynyl, lower
- 30 alkyl, lower alkoxy, formyl and phenyl;

002020" 56996460

or an isomer or pharmaceutically acceptable salt thereof.

32. Compound of Claim 34 wherein R^a is hydrido;
5 wherein R^b is selected from hydrido, and chloro;
wherein R^c is selected from chloro, methyl, tert-
butyl, methylthio, trifluoromethyl, difluoromethyl,
pentafluoromethyl, trifluoromethylsulfide,
trifluoromethoxy, cyano, substituted or
10 unsubstituted phenylcarbonyl, and substituted or
unsubstituted phenyl; wherein R^d is selected from
hydrido, methyl, tert-butyl, chloro; and wherein R^e is
selected from hydrido, chloro, thienyl,
hydroxyiminomethyl, substituted or unsubstituted
15 phenylethynyl, and substituted or unsubstituted
phenyl; or an isomer or pharmaceutically acceptable
salt thereof.

33. A method of treating a cyclooxygenase-2
20 mediated disorder in a subject, said method
comprising treating the subject having or susceptible
to said disorder with a therapeutically-effective
amount of a compound of Claims 1-31; or a
pharmaceutically-acceptable salt thereof.

25

34. The method of Claim 33 wherein the
cyclooxygenase-2 mediated disorder is inflammation.

35. The method of Claim 33 wherein the
30 cyclooxygenase-2 mediated disorder is arthritis.

36. The method of Claim 33 wherein the
cyclooxygenase-2 mediated disorder is pain.

35 37. The method of Claim 33 wherein the
cyclooxygenase-2 mediated disorder is fever.

002020" 56996460

38. A pharmaceutical composition comprising a therapeutically-effective amount of a compound, said compound selected from a family of compounds of Claims 1-31; or a pharmaceutically-acceptable salt thereof.
- 5

002020" 56996460